

Supplemental material:

Supplemental Methods

Search strategy and selection criteria; Genetic studies

Reference lists of articles identified from the primary search were additionally scanned for relevant articles, including previous meta-analyses and systematic reviews. For inclusion, genetic studies had to have more than 500 participants, involve unrelated subjects and be published as full length articles or letters in a peer reviewed journal. Authors of published studies were contacted (on at least 3 occasions) to obtain additional information on *CETP* genotypes and variables of interest, where unreported. The collaborative group for the genetic analysis was assembled by direct contact with principal investigators of any study known to the authors that involved more than 500 individuals that had previously reported at least one genetic finding, in any area, in a peer reviewed journal.

Twenty two studies were identified from the search and the published meta-analysis and seven were either reported after the previously published meta-analysis, or were contacted independently of the search as these studies were known to have published on genetic associations in lipids. Where data were duplicated in two publications, clarity was sought from the author, and limited tabular data were requested on the complete cohort. If there was no response, the larger of the data sets reported were included. Four unpublished cohorts were included.

Data Extraction

Randomised trials: Information was extracted on treatment regimen and comparator, pre- and post-treatment concentration of HDL-, LDL- and total cholesterol, triglycerides, apolipoproteins A-I (apoA-I) apolipoprotein B (apo-B), C-reactive protein (CRP), sodium, potassium, chloride, and bicarbonate, aldosterone, plasma creatinine and estimated glomerular filtration rate, systolic and diastolic blood pressure.

Genetic Studies: Information was obtained on study design, total number of participants and the number of individuals by genotype category, gender, ethnic origin and presence or absence of CHD at baseline. In addition, summary information on the following variables was obtained (where available) for each *CETP*-genotype group: *CETP* concentration, *CETP* activity, HDL-, LDL-, and total cholesterol, triglycerides, apoA-I and apoA-II, apoB, HDL sub-fractions 2 and 3, systolic and diastolic

blood pressure, blood glucose, CRP, urinary and plasma sodium, potassium and creatinine, body mass index, smoking status, age and treatment with anti-hypertensive medications or statins.

Statistical Analysis

For continuous traits, median values were assumed to be equal to mean values. If the standard deviation (SD) was not reported this was calculated from the standard error (SE) by multiplying the this by the square root of the sample size or from the inter-quartile range by dividing the width of the inter-quartile range by 1.349. For dose ranging studies, the SD was imputed from the largest study if unavailable. Where logged values were provided, these were back transformed and geometric means were used as means. Where cholesterol and triglycerides were reported in mg/ dl, values were converted to mmol/l by multiplying by 0.02586 and 0.01129 respectively. Similarly where glucose was reported as mg/dl values were converted to mmol/l by multiplying by 0.055, and creatinine was converted to $\mu\text{mol/L}$ by multiplying by 88. We estimated using a MAF of 0.48, a sample size of 14,147 individuals was required to be able to detect a difference of 0.5 mmHg SBP, with a power of 0.8 at a significance of 0.05 (calculated using online genetic power calculator “Quanto” (S40)).

Simulation of observed gene vs expected torcetrapib dose equivalent effects: Once a linear dose response was confirmed for available variables, the summary effect with most data from dose finding or randomized trials (60mg) on HDL-C was used for simulation studies. The effect of 60 mg torcetrapib on HDL was called β_{d1} (with standard error $sd1$), and for other i traits, similarly β_{di} (with standard error s_{di}). The simulation model then incorporated the variance in the effect estimates of the genotype and drug effects. This was done as follows: Firstly, a random draw (x_{d1}) of the 60mg drug effect on HDL cholesterol was taken from a normal distribution $N(\beta_{d1}, s_{d1})$ and of the effect on each other trait (random draw x_{di} from $N(\beta_{di}, s_{di})$, and of the gene effect on HDL cholesterol (draw x_{g1} from $N(\beta_{g1}, s_{g1})$). The gene effect (x_{g1}) was then expressed as a torcetrapib dose equivalent, calculated as $dose_g = 60 \times x_{g1} / x_{d1}$, from which the expected effect of this dose on trait i was estimated as $e_{gi} = x_{di} \times x_{g1} / x_{d1}$. Random draws were repeated 100000 times, generating a distribution of $dose_g$ from which its mean, 2.5th and 97.5th percentiles were used to estimate the mean (95% confidence interval) for the gene-dose equivalent. Similarly the mean of the distribution of expected effects on trait i was estimated (m_{egi}) as well as its standard error (s_{egi}) as $(97.5^{th} - 2.5^{th} \text{ percentiles}) / (2 \times 1.96)$. The observed gene

effect on trait i (β_{gi}) was compared to the expected effect of a comparable dose of torcetrapib ($m_{e_{gi}}$) using a Z- test, $z=(\beta_{gi}-m_{e_{gi}})/\sqrt{(s_{gi}^2+s_{e_{gi}}^2)}$.

Consistency between the CETP gene effects and equivalent torcetrapib dose

The 95% confidence intervals for the expected effect of a dose of torcetrapib comparable to the effect of genotype were obtained by simulation. To incorporate the uncertainty in the effect estimates, one hundred thousand replications were generated of the point estimates and standard errors of the 60 mg dose of torcetrapib. The values of the 2.5 and 97.5 centiles of the simulated distribution were used as the 95% confidence intervals. The simulation process was conducted separately for individuals homozygous for the B2 allele and then repeated for heterozygous individuals.

Supplemental Results

Seven studies with 21, 353 individuals homozygous for the rs5882 (I405V) allele also had higher concentrations of HDL cholesterol (0.04 mmol/L; 0.00, 0.09), although the effect is less marked than that of rs7082872 (Taq1B). Similarly there was no evidence of a link between the I405V variant and blood pressure.

Supplemental Tables:

Table S1: Effect of torcetrapib (60 mg) and *CETP* genotype on other continuous and demographic variables.

Differences between continuous traits are those reported at the end of the randomised trial end unless otherwise indicated. Differences in demographic variables for RCTs were those recorded at baseline. ** only ILLUMINATE contributed to analysis.

Comparison	RCTs, Torcetrapib 60 mg (Number of Individuals)	Summary Mean Difference/ Odds Ratio (95% CI)	P value	Genetic Studies, B1B2 vs B1B1, no of studies (Individuals)	Summary Mean Difference/ Odds Ratio (95%CI)	P Value	Genetic Studies, B2B2 vs B1B1, no of studies (Individuals)	Summary Mean Difference/ Odds Ratio (95%CI)	P Value
Continuous traits									
C-reactive protein (mg/L)	2 (17,007)	0.02 (-0.04, 0.08)	0.52	13 (34,826)	0.03 (-0.07, 0.13)	0.60	13 (22,049)	0.16 (0.04, 0.29)	0.01
Glucose (mmol/L)	NA	NA	NA	11 (32,608)	0.00 (-0.02, 0.02)	0.95	11 (20,497)	0.03 (0.00, 0.06)	0.09
Demographic and other variables									
Age	4 (17,911)	-0.48 (-1.29, 0.33)	0.25	19 (43,950)	0.00 (-0.20, 0.21)	0.98	19 (27,583)	0.18 (0.00, 0.35)	0.05
BMI (kg/m²)	4 (17,911)	-0.06 (-0.22, 0.1)	0.46	19 (40,212)	-0.07 (0.15, 0.02)	0.12	19 (25,249)	-0.01 (-0.12, 0.10)	0.85
Treatment with statin	4 (17,911)	All treated with Atorvastatin	All treated with Atorvastatin	7 (20,600)	1.02 (0.10, 10.74)	0.99	7 (13,005)	0.92 (0.09, 9.79)	0.95
Treatment anti-hypertensive medication	3 (2,844)	0.98 (0.07, 13.3)	0.99	7 (20,077)	0.99 (0.83,1.17)	0.87	10 (15,154)	0.98 (0.80, 1.21)	0.85
Hypertension at baseline	4 (17,911)	1.04 (0.05, 22.5)	0.99	NA	NA	NA	NA	NA	NA
Current/ former vs never smoked	NA	NA	NA	9 (23,420)	0.92 (0.82, 1.03)	0.16	9 (14,649)	0.90 (0.78,1.04)	0.15

Table S2. Studies contributing to the CETP analysis

Study	Year of publication	Country	Gender male (%)	Mean age (years)	Total sample (n)	Study Description	Baseline coronary heart disease (CHD)	HWE χ^2 for main genotype included	CETP SNP(s) typed
*Published studies with blood pressure, **studies that did not respond to data request									
REGRESS ^{S30*}	1998	Netherlands	100	56	807	Cases of CHD from RCT	CHD	0.33	rs708272
Corella D ^{S11**}	1999	Spain	45	36.6	514	Cross sectional	No CHD	1.18	rs708272
Rekyavik ^{S13**}	2000	Iceland	100	71	1,134	Prospective cohort study - cases of CHD only	CHD	4.45	rs708272
Brousseau M ^{S9**}	2002	United States	100	64	833	Cross Sectional	CHD	3.71	rs708272
Atherogene ^{S6**}	2003	Germany	75	64.4	1,211	Prospective cohort of CHD	CHD	1.98	rs1800775
CARE ^{S12*}	2004	Canada, United States	86	59.5	3,205	Cases from RCT	CHD	1.78	rs708272
PHS ^{S24**}	2005	United States	100	58.4	768	Prospective cohort	Mixed Population	1.01	rs708272
Marschang P ^{S31**}	2006	Austria	56	64.3	983	Prospective Cohort	CHD	0.06	rs708272
*Studies with unpublished data on blood pressure, *unpublished studies									
EARS ^{S16*}	1999	Estonia, Belgium, Denmark, Finland, Germany, Greece, Italy, Portugal, Spain, Switzerland, United Kingdom	100	23	794	Cross sectional	No CHD	0.00	rs708272, rs5882
OPERA ^{S18,S19,S20*}	2000	Finland	49	51.4	524	Prospective cohort	Mixed Population	0.02	rs708272, rs158477
Framingham Offspring ^{S29*}	2000	United States	48.4	51.3	2,916	Prospective cohort	Mixed Population	1.84	rs708272
Arca M ^{S4*}	2001	Italy	65.4	58.7	798	Case control, additional group of population controls	Mixed Population	0.88	rs708272
ECTIM ^{S10,S15,S22*}	2002	France, United Kingdom	78	55.5	2,540	Case control	Mixed Population	1.27	rs708272, rs5882, rs1800775, G-971A
NPFS ^{S26*}	2002	United Kingdom	100	55.9	2,589	Prospective Cohort	No CHD	3.4	rs708272
WOSCOPS ^{S14*}	2003	United Kingdom	100	57	1,604	RCT	Mixed Population	0.28	rs708272
ACCESS ^{S25*}	2005	United states	60	60.3	2,106	RCT	Mixed Population	0.01	72 SNPs including rs708282, rs5882, rs1800775
Sorli ^{S27*}	2006	Spain	31	45.5	549	Cross sectional	Mixed Population	0.99	rs708272
PREVEND ^{S8*}	2006	Netherlands	50.5	49.4	8,166	Prospective cohort	Mixed Population	1.56	rs708272, rs1800775, rs5882
SAPHIR ^{S28*}	2008	Austria	68	52.7	1,503	Prospective Cohort	No CHD	0.31	rs708272
Busseiton ^{S1*}	2007	Australia	45	49	1,574	Cross sectional	Mixed Population	1.05	rs708272, rs1800775, rs12149545
CUDAS ^{S1*}	2007	Australia	50	53	1,109	Cross sectional	Mixed Population	0.01	rs708272, rs1800775, rs12149546
CUPID ^{S1*}	2007	Australia	87	50	556	Cohort of patients presenting for coronary catheters	CHD	0.12	rs708272, rs1800775, rs12149547
Intermountain ^{S17*}	2007	United States	69	56	9,371	Cohort of patients presenting for coronary catheters	Mixed Population	1.22	32 SNPs including rs708282, rs5882, rs1800775
Health Professionals Study ^{S23*}	2007	United States	100	60	2,193	Prospective cohort study	No CHD	0.945	rs708272
Nurses Health Study ^{S5*}	2007	United States	0	57.4	1,291	Prospective cohort study	No CHD	1.3	rs708272
Rotterdam ^{S33*}	2007	Netherlands	41	68.8	6,421	Prospective cohort study	Mixed Population	0.48	rs5882, rs1800775
British Women's Heart and Health Study ^{S21**}	Unpublished	United Kingdom	0	68.9	3,570	Prospective cohort study	Mixed Population	2.58	rs708272
EPIC-Norfolk ^{S7**}	Unpublished	United Kingdom	63	65.3	2,114	Prospective cohort study	No CHD	2.2	rs1800775
Whitehall II ^{S3**}	Unpublished	United Kingdom	73	49.8	5,049	Prospective cohort study	Mixed Population	0.61	rs708272
ELSA ^{S2**}	Unpublished	United Kingdom	46	63.6	5,541	Prospective cohort study	Mixed Population	1.35	rs708272
Erasmus ^{S32**}	Unpublished	Netherlands	40.7	53.12	873	Prospective cohort study	Mixed Population	0.024	rs5882

Table S3. Traits included from studies evaluating *CETP* Taq1B and -629C>A genetic variant in European descent individuals (“0” for not included and “1” for included)

Study	HDL-C	LDL-C	Total Cholesterol	Triglycerides	HDL2	HDL3	ApoA1	ApoB	SBP	DBP	PP	Glucose	BMI	CRP	Electrolytes and renal measures	Use of antihypertensive medication	Smoking status	Use of Statin
REGRESS	1	1	1	1	0	0	0	0	1	1	0	0	1	0	0	1	1	1
Corella D	1	1	1	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0
Reykjavik	1	0	1	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0
Brousseau	1	1	1	1	1	1	1	1	0	0	0	0	1	0	0	0	0	0
Atherogene	1	1	1	1	0	0	1	1	0	0	0	0	0	0	0	0	0	0
CARE	1	1	1	1	0	0	0	0	1	1	0	0	1	0	0	0	0	0
PHS	1	0	1	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0
Marschang P	1	1	1	1	0	0	0	0	0	0	0	0	1	0	0	0	0	0
EARS	1	1	1	1	0	0	1	1	1	1	1	0	1	1	1	0	1	0
OPERA	1	1	1	1	0	0	0	0	1	1	0	0	1	0	1	0	0	0
Framingham	1	1	1	1	1	1	1	1	1	1	1	0	1	1	0	0	0	0
Arca M	1	1	1	1	0	0	0	0	1	1	0	0	0	0	1	0	0	0
ECTIM	1	1	1	1	0	0	1	1	1	1	1	0	0	1	0	0	1	0
NPHS II	1	1	1	1	0	0	1	1	1	1	1	1	1	1	1	0	1	0
WOSCOPS	1	1	1	1	0	0	1	1	1	1	1	0	1	1	1	1	1	1
ACCESS	1	1	1	1	0	0	0	0	1	1	0	0	0	0	0	0	0	0
Sorli	1	1	1	1	0	0	0	0	1	1	0	0	0	0	0	0	0	0
PREVEND	1	1	1	1	0	0	1	1	1	1	1	1	1	1	1	1	1	1
SAPHIR	1	1	1	1	0	0	1	1	1	1	0	1	1	1	1	1	1	0
Busselton	1	1	1	1	0	0	0	0	1	1	0	0	1	0	0	0	0	0
CUDAS	1	1	1	1	0	0	0	0	1	1	0	1	1	1	0	0	0	1
CUPID	1	1	1	1	0	0	0	0	1	1	0	1	1	1	1	0	0	1
Intermountain Health	1	1	1	1	0	0	0	0	1	1	1	1	1	1	1	1	1	1
Professionals Study	1	1	1	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0
Nurses Health Study	1	1	1	1	0	0	0	0	0	0	0	0	0	0	0	0	0	0
BWHHS	1	1	1	1	0	0	0	0	1	1	1	1	1	1	1	1	1	1
EPIC	1	1	1	1	0	0	1	1	1	1	1	1	1	1	1	1	1	0
Rotterdam	1	0	1	0	0	0	0	0	1	1	0	0	0	0	1	0	0	0
Whitehall II	1	1	1	1	0	0	1	1	1	1	1	1	1	1	0	0	1	0
ELSA	1	1	1	1	0	0	0	0	1	1	1	1	0	1	0	1	1	0

Supplemental Figures and Figure Legends:

Figure S1 LD structure of the *CETP* gene. r^2 values are given from the ACCESS study. The main SNPs evaluated in this study were Taq1B (rs708272) and -629C>A (rs1800775) which are in LD ($r^2=0.73$). SNPs contributing to variance in HDL cholesterol identified from genome wide association scans are also shown (rs12596776, rs2217332, rs3764261, rs1800775, rs711752, rs1864163, rs7205804, rs5880, rs5882, rs1800777, rs1566439). Data provided by J F Thompson, ACCESS study¹⁶

Figure S2a and S2b Standardised mean differences in lipid and lipoproteins between individuals homozygous for *CETP* variants in populations studies (a) and those receiving torcetrapib 60mg (b) daily as compared to placebo in clinical trials

Figures S3a-c Effect of *CETP* genotype on (a) CETP concentration, (b) CETP activity and (c) HDL cholesterol concentration. Forest plots indicate weighted mean difference and 95% confidence intervals. Results are stratified by ancestral origin, study size, and prevalent coronary heart disease, gender and polymorphism typed. (*the B1B1 genotype grouped is used as the reference group throughout)

Figure S4: Association between *CETP* genotype (B2B2 vs B1B1) and HDL-cholesterol level stratified by systolic blood pressure (SBP), diastolic blood pressure (DBP), pulse pressure (PP) and LDL-Cholesterol. Data from 6 studies, (12,983 individuals)

Figure S5a-b Effect of *CETP* genotype on (a) systolic and (b) diastolic blood pressure in populations of European descent only. Forest plots show weighted mean difference and 95% confidence intervals. Results are stratified by study size, prevalent coronary heart disease, gender, polymorphism typed, and strata of LDL cholesterol. (The B1B1 genotype is used as the reference group, see text for details)

Figure S1

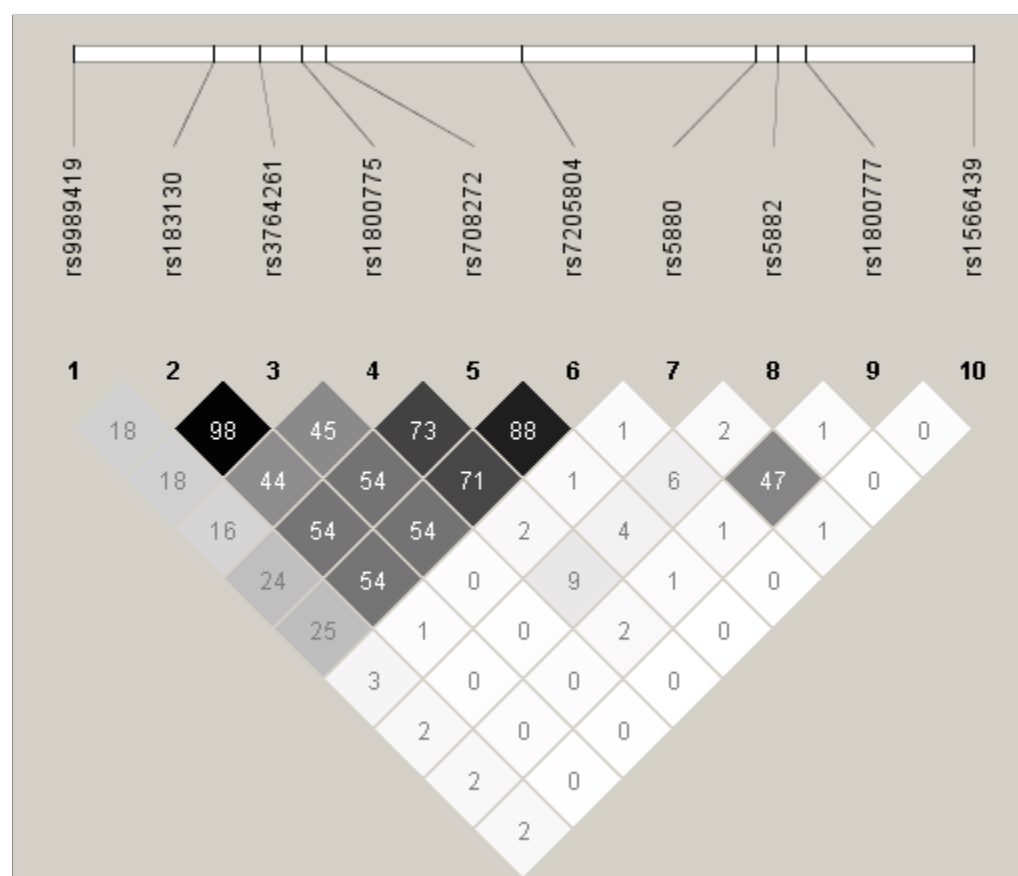


Figure S2a

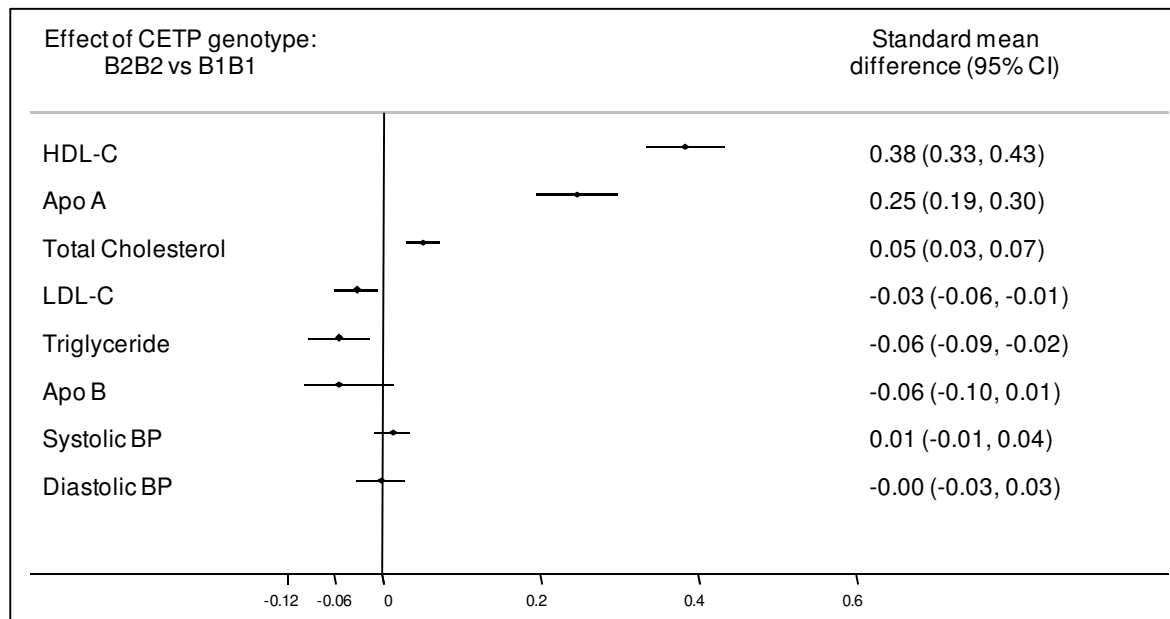


Figure S2b

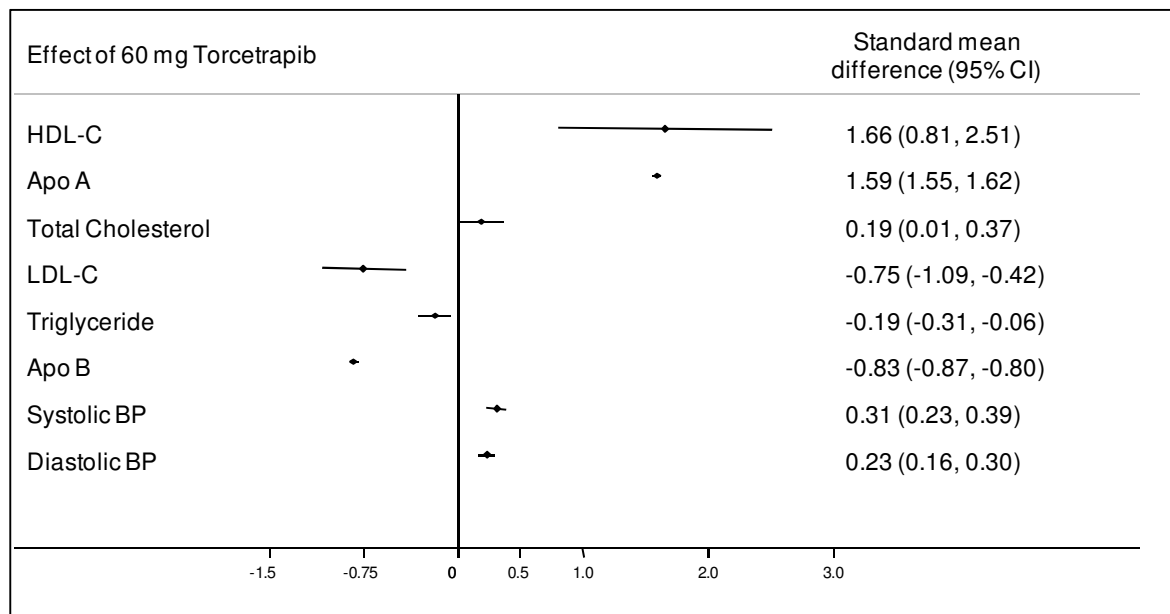


Figure S3a

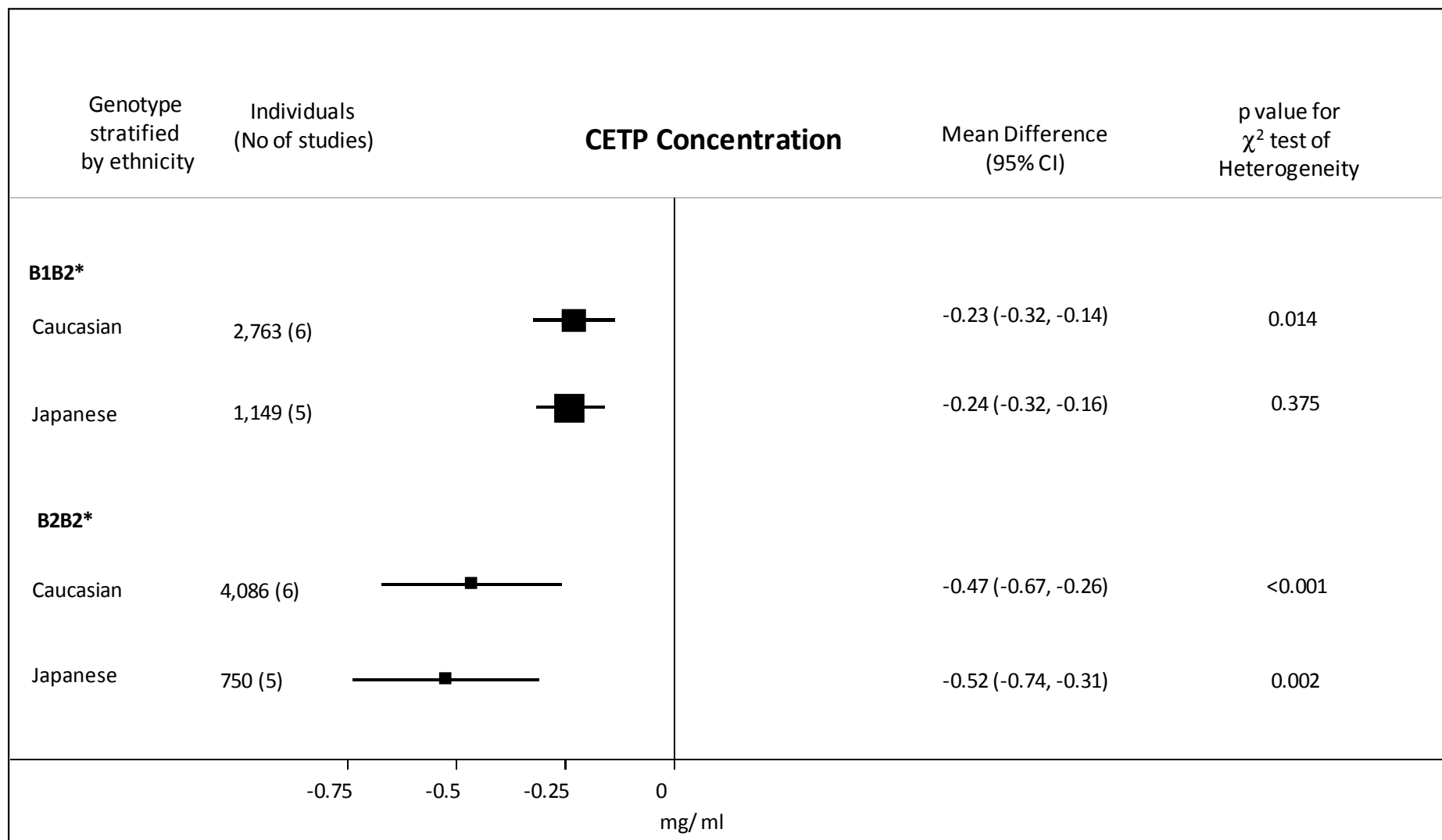


Figure S3b

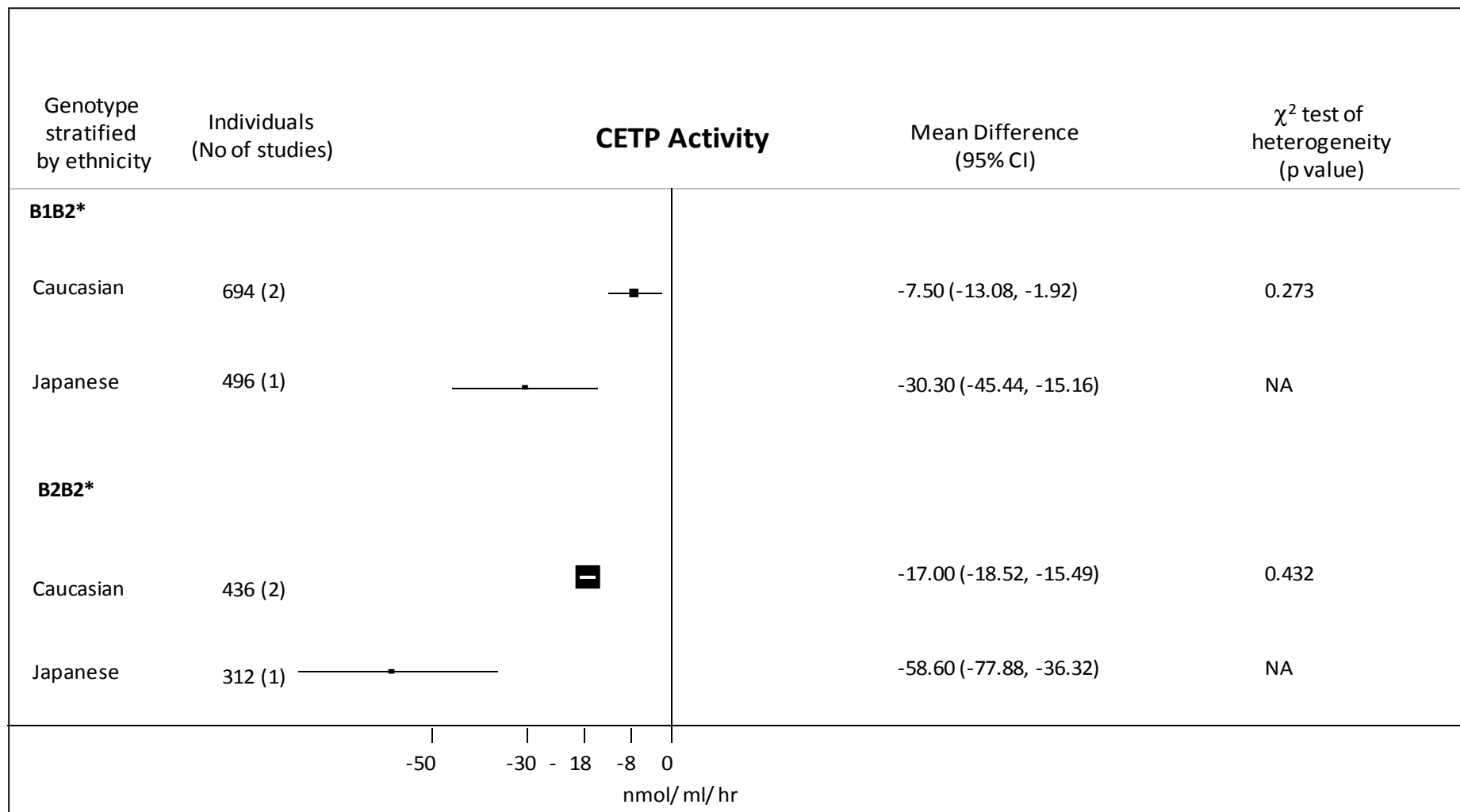


Figure S3c

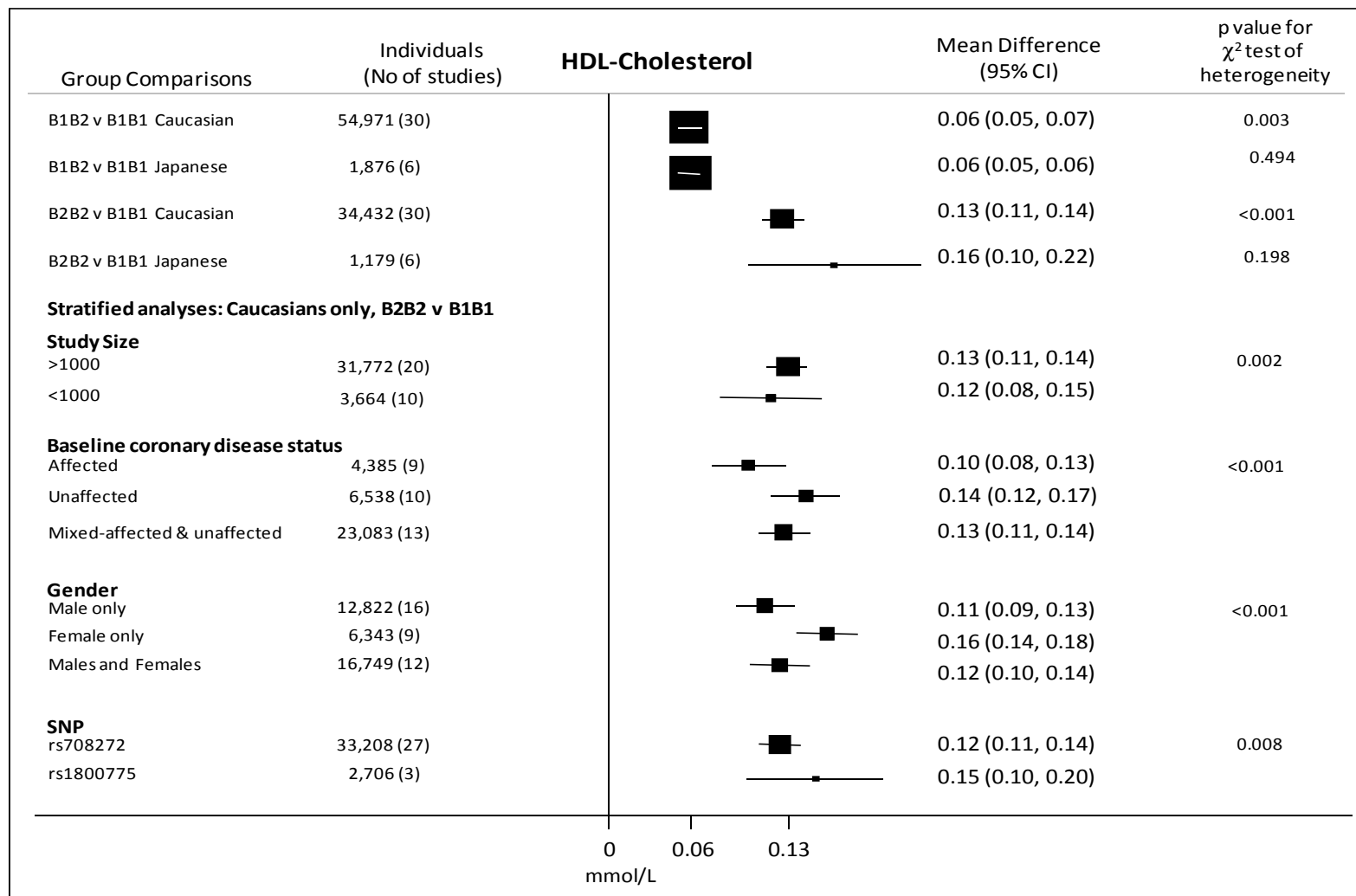


Figure S4

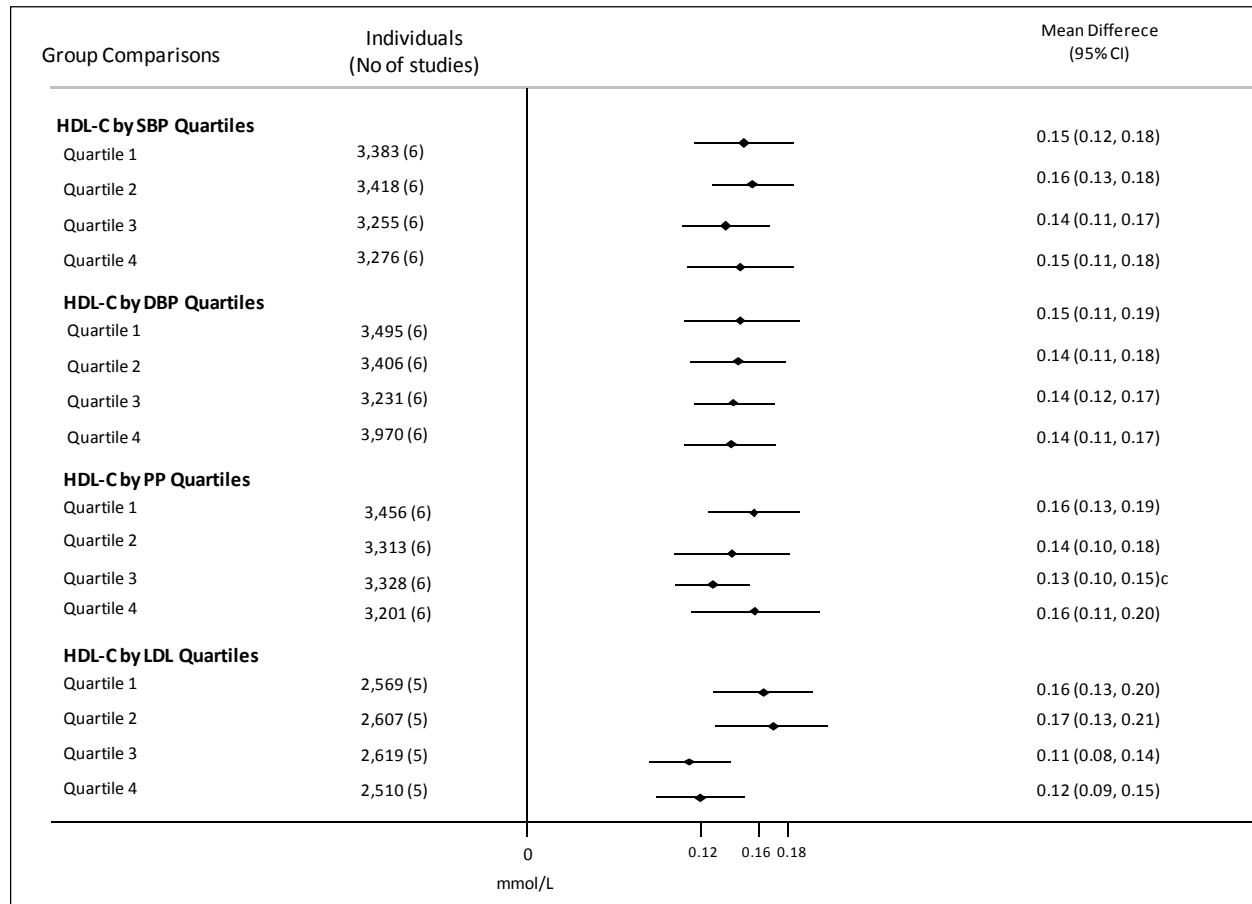


Figure S5a

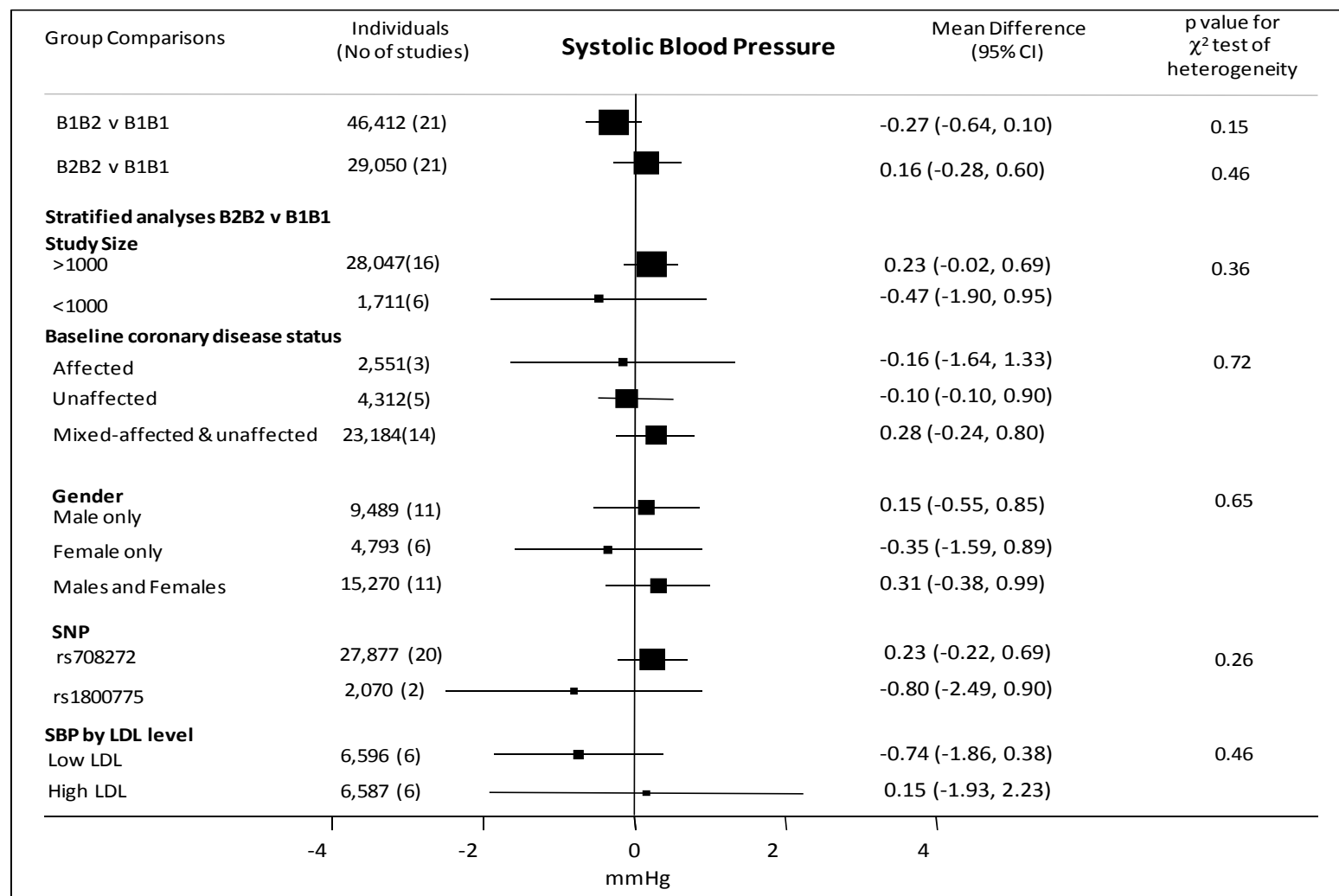
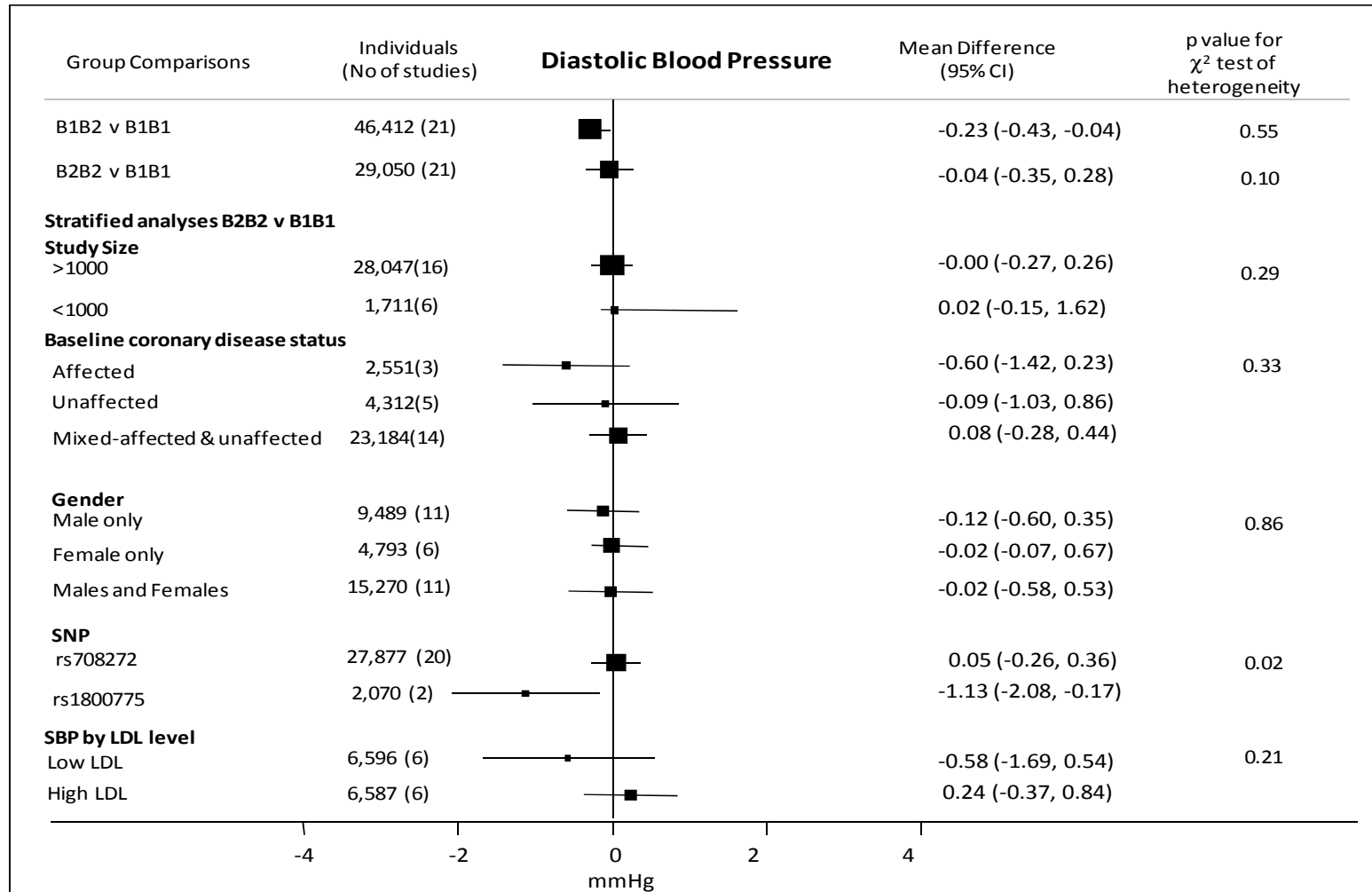


Figure S5b



Supplementary References:

- S1. McCaskie PA, Beilby JP, Chapman CM, et al. Cholesteryl ester transfer protein gene haplotypes, plasma high-density lipoprotein levels and the risk of coronary heart disease. *Hum Genet* 2007;121:401-11.
- S2. Marmot M, Banks J, Blundell R, Lessoff C, Nazroo J. Health, wealth and lifestyles of the older population in England: THE 2002 ENGLISH LONGITUDINAL STUDY OF AGEING: Institute of Fiscal Studies; 2002.
- S3. Marmot M, Brunner E. Cohort Profile: the Whitehall II study. *Int J Epidemiol* 2005;34:251-6.
- S4. Arca M, Montali A, Ombres D, et al. Lack of association of the common TaqIB polymorphism in the cholesteryl ester transfer protein gene with angiographically assessed coronary atherosclerosis. *Clin Genet* 2001;60:374-80.
- S5. Asselbergs FW, Pai JK, Rexrode KM, Hunter DJ, Rimm EB. Effects of lymphotoxin-alpha gene and galectin-2 gene polymorphisms on inflammatory biomarkers, cellular adhesion molecules and risk of coronary heart disease. *Clin Sci (Lond)* 2007;112:291-8.
- S6. Blankenberg S, Rupprecht HJ, Bickel C, et al. Common genetic variation of the cholesteryl ester transfer protein gene strongly predicts future cardiovascular death in patients with coronary artery disease. *J Am Coll Cardiol* 2003;41:1983-9.
- S7. Boekholdt SM, Kuivenhoven JA, Wareham NJ, et al. Plasma levels of cholesteryl ester transfer protein and the risk of future coronary artery disease in apparently healthy men and women: the prospective EPIC (European Prospective Investigation into Cancer and nutrition)-Norfolk population study. *Circulation* 2004;110:1418-23.
- S8. Borggreve SE, Hillege HL, Wolffenbuttel BH, et al. The effect of cholesteryl ester transfer protein -629C>A promoter polymorphism on high-density lipoprotein cholesterol is dependent on serum triglycerides. *J Clin Endocrinol Metab* 2005;90:4198-204.
- S9. Brousseau ME, O'Connor JJ, Jr., Ordovas JM, et al. Cholesteryl ester transfer protein TaqI B2B2 genotype is associated with higher HDL cholesterol levels and lower risk of coronary heart disease end points in men with HDL deficiency: Veterans Affairs HDL Cholesterol Intervention Trial. *Arterioscler Thromb Vasc Biol* 2002;22:1148-54.
- S10. Corbex M, Poirier O, Fumeron F, et al. Extensive association analysis between the CETP gene and coronary heart disease phenotypes reveals several putative functional polymorphisms and gene-environment interaction. *Genet Epidemiol* 2000;19:64-80.

- S11. Corella D, Saiz C, Guillen M, et al. Association of TaqIB polymorphism in the cholesteryl ester transfer protein gene with plasma lipid levels in a healthy Spanish population. *Atherosclerosis* 2000;152:367-76.
- S12. de Grooth GJ, Zerba KE, Huang SP, et al. The cholesteryl ester transfer protein (CETP) TaqIB polymorphism in the cholesterol and recurrent events study: no interaction with the response to pravastatin therapy and no effects on cardiovascular outcome: a prospective analysis of the CETP TaqIB polymorphism on cardiovascular outcome and interaction with cholesterol-lowering therapy. *J Am Coll Cardiol* 2004;43:854-7.
- S13. Eiriksdottir G, Bolla MK, Thorsson B, Sigurdsson G, Humphries SE, Gudnason V. The -629C>A polymorphism in the CETP gene does not explain the association of TaqIB polymorphism with risk and age of myocardial infarction in Icelandic men. *Atherosclerosis* 2001;159:187-92.
- S14. Freeman DJ, Samani NJ, Wilson V, et al. A polymorphism of the cholesteryl ester transfer protein gene predicts cardiovascular events in non-smokers in the West of Scotland Coronary Prevention Study. *Eur Heart J* 2003;24:1833-42.
- S15. Fumeron F, Betoulle D, Luc G, et al. Alcohol intake modulates the effect of a polymorphism of the cholesteryl ester transfer protein gene on plasma high density lipoprotein and the risk of myocardial infarction. *J Clin Invest* 1995;96:1664-71.
- S16. Gudnason V, Kakko S, Nicaud V, et al. Cholesteryl ester transfer protein gene effect on CETP activity and plasma high-density lipoprotein in European populations. The EARS Group. *Eur J Clin Invest* 1999;29:116-28.
- S17. Horne BD, Camp NJ, Anderson JL, et al. Multiple less common genetic variants explain the association of the cholesteryl ester transfer protein gene with coronary artery disease. *J Am Coll Cardiol* 2007;49:2053-60.
- S18. Kakko S, Tamminen M, Paivansalo M, et al. Cholesteryl ester transfer protein gene polymorphisms are associated with carotid atherosclerosis in men. *Eur J Clin Invest* 2000;30:18-25.
- S19. Kakko S, Tamminen M, Paivansalo M, et al. Variation at the cholesteryl ester transfer protein gene in relation to plasma high density lipoproteins cholesterol levels and carotid intima-media thickness. *Eur J Clin Invest* 2001;31:593-602.
- S20. Kauma H, Savolainen MJ, Heikkila R, et al. Sex difference in the regulation of plasma high density lipoprotein cholesterol by genetic and environmental factors. *Hum Genet* 1996;97:156-62.
- S21. Lawlor DA, Day IN, Gaunt TR, et al. The association of the PON1 Q192R polymorphism with coronary heart disease: findings from the British Women's Heart and Health cohort study and a meta-analysis. *BMC Genet* 2004;5:17.

- S22. Le Goff W, Guerin M, Nicaud V, et al. A novel cholesteryl ester transfer protein promoter polymorphism (-971G/A) associated with plasma high-density lipoprotein cholesterol levels. Interaction with the TaqIB and -629C/A polymorphisms. *Atherosclerosis* 2002;161:269-79.
- S23. Li TY, Zhang C, Asselbergs FW, et al. Interaction between dietary fat intake and the cholesterol ester transfer protein TaqIB polymorphism in relation to HDL-cholesterol concentrations among US diabetic men. *Am J Clin Nutr* 2007;86:1524-9.
- S24. Liu S, Schmitz C, Stampfer MJ, et al. A prospective study of TaqIB polymorphism in the gene coding for cholesteryl ester transfer protein and risk of myocardial infarction in middle-aged men. *Atherosclerosis* 2002;161:469-74.
- S25. Lloyd DB, Lira ME, Wood LS, et al. Cholesteryl ester transfer protein variants have differential stability but uniform inhibition by torcetrapib. *J Biol Chem* 2005;280:14918-22.
- S26. Talmud PJ, Hawe E, Robertson K, Miller GJ, Miller NE, Humphries SE. Genetic and environmental determinants of plasma high density lipoprotein cholesterol and apolipoprotein AI concentrations in healthy middle-aged men. *Ann Hum Genet* 2002;66:111-24.
- S27. Sorli JV, Corella D, Frances F, et al. The effect of the APOE polymorphism on HDL-C concentrations depends on the cholesterol ester transfer protein gene variation in a Southern European population. *Clin Chim Acta* 2006;366:196-203.
- S28. Sandhofer A, Tatarczyk T, Laimer M, et al. The Taq1B-variant in the Cholesteryl Ester-Transfer Protein Gene and the Risk of Metabolic Syndrome. *Obesity* (Silver Spring) 2008.; Apr16(4):919-22
- S29. Ordovas JM, Cupples LA, Corella D, et al. Association of cholesteryl ester transfer protein-TaqIB polymorphism with variations in lipoprotein subclasses and coronary heart disease risk: the Framingham study. *Arterioscler Thromb Vasc Biol* 2000;20:1323-9.
- S30. Kuivenhoven JA, Jukema JW, Zwinderman AH, et al. The role of a common variant of the cholesteryl ester transfer protein gene in the progression of coronary atherosclerosis. The Regression Growth Evaluation Statin Study Group. *N Engl J Med* 1998;338:86-93.
- S31. Marschang P, Sandhofer A, Ritsch A, Fiser I, Kvas E, Patsch JR. Plasma cholesteryl ester transfer protein concentrations predict cardiovascular events in patients with coronary artery disease treated with pravastatin. *J Intern Med* 2006;260:151-9.
- S32. Isaacs A, Sayed-Tabatabaei FA, Aulchenko YS, et al. Heritabilities, apolipoprotein E, and effects of inbreeding on plasma lipids in a genetically isolated population: the Erasmus Rucphen Family Study. *Eur J Epidemiol* 2007;22:99-105.
- S33. Isaacs A, Sayed-Tabatabaei FA, Hofman A, et al. The cholesteryl ester transfer protein I405V polymorphism is associated with increased high-density lipoprotein levels and decreased risk of myocardial infarction: the Rotterdam Study. *Eur J Cardiovasc Prev Rehabil* 2007;14:419-21.

- S34. Arai H, Yamamoto A, Matsuzawa Y, et al. Polymorphisms in four genes related to triglyceride and HDL-cholesterol levels in the general Japanese population in 2000. *J Atheroscler Thromb* 2005;12:240-50.
- S35. Goto A, Sasai K, Suzuki S, et al. Cholesteryl ester transfer protein and atherosclerosis in Japanese subjects: a study based on coronary angiography. *Atherosclerosis* 2001;159:153-63.
- S36. Ikewaki K, Mabuchi H, Teramoto T, et al. Association of cholesteryl ester transfer protein activity and TaqIB polymorphism with lipoprotein variations in Japanese subjects. *Metabolism* 2003;52:1564-70.
- S37. Kawasaki I, Tahara H, Emoto M, Shoji T, Nishizawa Y. Relationship between TaqIB cholesteryl ester transfer protein gene polymorphism and macrovascular complications in Japanese patients with type 2 diabetes. *Diabetes* 2002;51:871-4.
- S38. Meguro S, Takei I, Murata M, et al. Cholesteryl ester transfer protein polymorphism associated with macroangiopathy in Japanese patients with type 2 diabetes. *Atherosclerosis* 2001;156:151-6.
- S39. Okumura K, Matsui H, Kamiya H, Saburi Y, Hayashi K, Hayakawa T. Differential effect of two common polymorphisms in the cholesteryl ester transfer protein gene on low-density lipoprotein particle size. *Atherosclerosis* 2002;161:425-31.
- S40. Gauderman WJ, Morrison JM. QUANTO 1.1: A computer program for power and sample size calculations for genetic-epidemiology studies, <http://hydra.usc.edu/gxe>, 2006